YATIONAL SEARCH REPORT

Application No PCT/EP2005/000609

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/37 G01N33/573 A61K38/00 A61K39/00 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12Q G01N A61K Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, BIOSIS, EMBASE, CHEM ABS Data, PAJ, WPI Data C. DOCUMENTS CONSIDERED TO BE RELEVANT Category ° Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Х WO 01/46443 A (INCYTE GENOMICS, INC; YANG, 1 - 26JUNMING; BAUGHN, MARIAH, R; BURFORD, NEIL;) 28 June 2001 (2001-06-28) page 43, line 28 page 40, line 12 - line 22 page 55, line 22 - line 28 page 44, line 25 - page 47, line 22 page 47, line 35 - page 48, line 26 claims 16,25-27 sequence 9 -/--Further documents are listed in the continuation of box C. Patent family members are listed in annex. " Special categories of cited documents: T later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance invention "E" earlier document but published on or after the International "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken-alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "O" document referring to an oral disclosure, use, exhibition or "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 2 2 08. **2005** 14 June 2005 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

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Form PCT/ISA/210 (second sheet) (January 2004)

INTERNATIONAL SEARCH REPORT

Internation Application No
PCT/EP2005/000609

Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	PCT/EP2005/000609	
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
X	MINNASCH PETRA ET AL: "Demonstration of puromycin-sensitive alanyl aminopeptidase in Alzheimer disease brain." LEGAL MEDICINE (TOKYO, JAPAN) MAR 2003, vol. 5 Suppl 1, March 2003 (2003-03), pages S285-S287, XP001206793 ISSN: 1344-6223 the whole document	1-26	
X .	TOBLER A R ET AL: "Cloning of the human puromycin-sensitive aminopeptidase and evidence for expression in neurons" JOURNAL OF NEUROCHEMISTRY, vol. 68, no. 3, 1997, pages 889-897, XP002331835 ISSN: 0022-3042 the whole document	1-26	
(HUBER G ET AL: "CDNA CLONING AND MOLECULAR CHARACTERIZATION OF HUMAN BRAIN METALLOPROTEASE MP100: A BETA-SECRETASE CANDIDATE?" JOURNAL OF NEUROCHEMISTRY, NEW YORK, NY, US, vol. 72, no. 3, 1999, pages 1215-1223, XP000911070 ISSN: 0022-3042	1-26	
	SCHÖNLEIN C ET AL: "Purification and characterization of a novel metalloprotease from human brain with the ability to cleave substrates derived from the N-terminus of beta-amyloid protein." BIOCHEMICAL AND BIOPHYSICAL RESEARCH COMMUNICATIONS. 30 MAY 1994, vol. 201, no. 1, 30 May 1994 (1994-05-30), pages 45-53, XP002331836 ISSN: 0006-291X the whole document	1-26	
	MARTINEZ JOSE M ET AL: "Aminopeptidase activities in breast cancer tissue" CLINICAL CHEMISTRY, vol. 45, no. 10, October 1999 (1999-10), pages 1797-1802, XP002331837 ISSN: 0009-9147 the whole document	1-26	
	WO 97/38114 A (NOVARTIS AG; FONTANA, ADRIANO; CONSTAM, DANIEL, BEAT; TOBLER, ANDREAS,) 16 October 1997 (1997-10-16) p.51, last paragraph to p.53, paragraph 3 (included).sequence 6	1-26	



interpolation No. PCT/EP2005/000609

Box II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: Although claim 26 is directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
see additional sheet
As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
A. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-26 (all partially)
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-26 (all partially)

Methods of screening for therapeutic agents useful in the treatment of Alzheimer disease. A method for diagnosing Alzheimer disease. Pharmaceutical compositions for the treatment of Alzheimer disease. Use of regulators of a NPEPPS for the preparation of a pharmaceutical composition for the treatment of Alzheimer disease. A method for the preparation of a pharmaceutical composition useful for the treatment of Alzheimer disease. Use of a regulator of NPEPPS for the regulation of NPEPPS activity in a mammal having Alzheimer disease (see p. 41).

2. claims: 1-26 (all partially)

Same subject-matter as invention 1 but relating to thyroid tumor, to ileum tumor, to lung tumor, to breast tumor and to kidney tumor (see p. 58).

3. claims: 1-26 (all partially)

Same subject-matter as invention 1 but relating to lung chronic obstructive pulmonary disease (COPD) (inflammatory and respiratory disease) (see p. 58 and 59).

4. claims: 1-26 (all partially)

Same subject-matter as invention 1 but relating to prostate benign prostatic hyperplasia (BPH) (urological disease) (see p. 58).

INTERNATIONAL SEARCH REPORT

mation on patent family members

Internati Application No
PCT/EP2005/000609

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 0146443	A	28-06-2001	AU CA EP JP WO	2285701 A 2394789 A1 1240335 A2 2004500812 T 0146443 A2	03-07-2001 28-06-2001 18-09-2002 15-01-2004 28-06-2001
WO 9738114	Α	16-10-1997	WO AU	9738114 A1 5686896 A	16-10-1997 29-10-1997